

Nucleotide sequence of a rabbit genomic DNA encoding mature endothelin-3

Shoichi Ohkubo, Yasuaki Itoh, Chiharu Kimura, Haruo Onda and Masahiko Fujino

Tsukuba Research Laboratories, Takeda Chemical Industries Ltd, Wadai 7, Tsukuba, Ibaraki 300-42, Japan

Submitted November 29, 1989

EMBL accession no. X15724

Endothelin, a potent vasoconstrictor peptide, was isolated and its cDNAs from human and porcine libraries were cloned (1, 2). Analysis of human genomic DNA suggested the existence of an ET-family (ET-1, ET-2, ET-3) (3). Using ET-3 specific antibody, we have recently revealed the ET-3 expression in the brain, intestine and placenta of rat (4). We have also isolated a human cDNA clone for ET-3 (5). To examine the species specificity of ET-3 genes, we have examined a rabbit ET-3 gene and present here its partial DNA sequence.

The rabbit genomic library (EMBL-3) was constructed from liver chromosomal DNA and screened (6) with a 45-nucleotide synthetic DNA probe corresponding to mature ET-1. The predicted amino acid sequence of rabbit ET-3 (boxed in the figure) is identical to human and rat sequences (3,7), and the prepro-ET-3 region is highly conserved, suggesting its important role in animals. Compared with human ET-3 gene (unpublished data) or its cDNA (5,9), this sequence includes the entire exon

containing mature region and its 3'-exon (probably exon 1). The predicted splice sites and junction consensus sequences in the rabbit gene are indicated by arrows and underlining, respectively. The first methionine at position 317 is thought to be the translation initiation site. The length of the 3'-portion of rabbit prepro-ET-3 is the same as the human type except for the 24bp repeat sequence which exists in both the human genome and cDNA but not in the rabbit genome.

REFERENCES

1. Yanagisawa, M. et al. (1988) *Nature* **332**, 411-415.
2. Ito, Y. et al. (1988) *FEBS Lett.* **231**, 440-444.
3. Inoue, A. et al. (1989) *Proc. Natl. Acad. Sci. USA* **86**, 2863-2867.
4. Matsumoto, H. et al. (1989) *Biochim. Biophys. Res. Commun.* **164**, 74-80.
5. Onda, H. et al. *FEBS Lett.* in press.
6. Maniatis, T. et al. (1982) Molecular Cloning (Cold Spring Harbor, NY).
7. Yanagisawa, M. et al. (1988) *Proc. Natl. Acad. Sci. USA* **85**, 6964-6967.
8. Bloch, K.D. et al. (1989) *J. Biol. Chem.* **264**, 18156-18161.

TGTGTTGGCGCGACCCCGGGCGCCCTGGTCAAAGGCCCGCGGGCAGCTCCAGCCCCCTCCGGGGGGCGGGAGGCAGGGGG	90
GTGGTGGAGGCCAGAAAAGCCCGAGCCCACAGCCGGGAGTCCTCTGGCGGGGATGGCGACGGCGCGCTGAAAGTTGGTACCGCCGCAAC	180
CCAACTGCGCGCTGCAGCCAGGACCGAGCGAGCCAGGGAGCCGGCGGGCTCGAACCCCCACGGCGAGCCCGCGGGCGCTGTACCTGG	270
CCACCCAGCGGGGACCTGCGCCCGGTGCTCCCGCCCTGATCCGGGTCATGGAGCCGGCTGTGGATCCTTTGGGGCTCACAGTG	360
M E P G L W I L L G L T V	13
ACCGCCGCCG <u>CAGTA</u> AGCGGCCCGGGCGCGCGCTGTCGGCGCGAGCGCACACAAAAGGACCTGGCGGGGAGGTGGCGCGTCGC	450
T A A A	17
GGGGAGGGCCCGCACCCCTGGAGGGCGCTGGCGGGGCGACAGCTCAGCGCAGGGCCTGCACGTGTGGCTATGGGGCTG	540
GTCAGCTACTGGTCAGTGTCTCGAAGGCTCTGCAGACTGCAGAACTGCTAGCCAAGTTTCAGTGGCCGAGCAGAGTGCCTGAA	630
GTTTCAGGGAGTTAGATGGCTTGAGGCTCTGCAGGGAGTGGCTGGAGGAAGCTTGCAAGCGCTTCACACGGCATGCACGGGCTTGG	720
AACATTTCTGAAACTGTGTTAGGGCTGTGCTGGCTGGAGGCTTGGAGGCTTGGAGGATGTGAGACAATTGGAGATAACTTGC	810
AGGAGTTGCTGCCCGCCCGAACCCAGGTGGCTGCCGGGTGGCGCTGAGGGCTCTGCTCTGGCTCAGGAG <u>CCCTGGTCTTGG</u>	900
<u>CTCCCTGAGGATTCTGCGCTTGCCCCAGACTGGGGGTGCTGCAGGACAGCGTGCCTGGGGGGGGGGGGGGGGGGGGGGGGGGGGGG</u>	990
G F V P C P Q T G G A G R T S V P R A P R V A G S E G	44
GAAGTGTGAAGACTCTGGCCAGCCCTAGAAGGCAGACTGTGGCCCCAACGGCAGGGCAAGGGCCAGGCCCTGGAGGCCCTGGCGGGGG	1080
D C E D S V A S P R R Q T V A P T A G K G P S P G S P G R G	74
CAGGGGGGGAGGGGAGCCGGGGCACCGCCGTGTGGAGCAGTGGCCACCTGCTCACCTAACAAAGACAAAGAGTGGCTACTACTGCCAC	1170
Q A A E G D P G H R R R C T C F T Y K D K E C V Y Y C H	104
CTGGACATCTGGATCAACACTCCGGAGTGAGTCAGGCCAGCCCCCCTGCCTCACGTGCCCCGTCAGGGGGCTGATGCCACC	1260
L D I I W I N T P (*)	113
CCTCAGCCCTGCGGGTCTCCGCCAGCCGGTCCCGCTGCAGGTCATGTGAGCCGCTGTCCTCTCCCTGGCCCTGGCCCCGGG	1350
GAAACCCAATCTGCCCTGTGTGGCGGAGAGGCCCTCCAGGAGCAGTGAGCTGGAGAGAGGATTAGACACCGAGACGCCAGGAGTT	1440
GAGGTGGAGGAAAGCCCGAAAGTGTGCTGCCAGCCAGGGCTGAGTGTGCTTGACCTGCTGACCTCCAGGCTGCCAGGAGTT	1530
TAGAATT	1538